<110> Armour, Kathryn L Clark, Michael R Williamson, Lorna M

<120> Binding Molecules Derived From Immunoglobulins Which Do Not Trigger Complement Mediated Lysis

<130> 620-117

<140> US 09/674,857

<141> 2000-11-07

<150> PCT/GB99/01441

<151> 1999-05-07

<150> GB 9809951.8

<151> 1998-05-08

<160> 27

<170> PatentIn Ver. 2.1

<210> 1

<211> 109

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Mutated
 antibody

<400> 1

Ala Pro Pro Val Ala Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro 1 5 10 15

Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val 20 25 30

Val Asp Val Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr Val
35 40 45

Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln 50 55 60

Tyr Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln
65 70 75 80

Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Gly 85 90 95

Leu Pro Ser Ser Ile Glu Lys Thr Ile Ser Lys Ala Lys 100 105

<210> 2

<211> 109

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Mutated
 antibody

<400> 2

Ala Pro Pro Val Ala Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro 1 5 10 15

Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val 20 25 30

Val Asp Val Ser His Glu Asp Pro Glu Val Gln Phe Asn Trp Tyr Val 35 40 45

Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln 50 55 60

Phe Asn Ser Thr Phe Arg Val Val Ser Val Leu Thr Val Val His Gln 65 70 75 80

Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Gly
85 90 95

Leu Pro Ser Ser Ile Glu Lys Thr Ile Ser Lys Thr Lys
100 105

<210> 3

<211> 110

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Mutated
 antibody

<400> 3

Ala Pro Pro Val Ala Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys
1 5 10 15

Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val 20 25 30

Val Val Asp Val Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr 35 40 45

Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu 50 55 60

Gln Tyr Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His 65 70 75 80 Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys 85 90 95

Gly Leu Pro Ser Ser Ile Glu Lys Thr Ile Ser Lys Ala Lys 100 105 110

<210> 4

<211> 110

<212> PRT

<213> Homo sapiens

<400> 4

Ala Pro Glu Leu Leu Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys
1 5 10 15

Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val 20 25 30

Val Val Asp Val Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr 35 40 45

Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu
50 55 60

Gln Tyr Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His 65 70 75 80

Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys 85 90 95

Ala Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Ala Lys 100 105 110

<210> 5

<211> 109

<212> PRT

<213> Homo sapiens

<400> 5

Ala Pro Pro Val Ala Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro 1 5 10 15

Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val 20 25 30

Val Asp Val Ser His Glu Asp Pro Glu Val Gln Phe Asn Trp Tyr Val 35 40 45

Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln 50 55 60

Phe Asn Ser Thr Phe Arg Val Val Ser Val Leu Thr Val Val His Gln 65 70 75 80

Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Gly 85 90 95

Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Thr Lys 100 105

<210> 6

<211> 110

<212> PRT

<213> Homo sapiens

<400> 6

Ala Pro Glu Leu Leu Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys
1 5 10 15

Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val 20 25 30

Val Val Asp Val Ser His Glu Asp Pro Glu Val Gln Phe Lys Trp Tyr
35 40 45

Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu 50 55 60

Gln Tyr Asn Ser Thr Phe Arg Val Val Ser Val Leu Thr Val Leu His 65 70 75 80

Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys 85 90 95

Ala Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Thr Lys 100 105 110

<210> 7

<211> 110

<212> PRT

<213> Homo sapiens

<400> 7

Ala Pro Glu Phe Leu Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys
1 5 10 15

Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val

Val Val Asp Val Ser Gln Glu Asp Pro Glu Val Gln Phe Asn Trp Tyr 35 40 45

Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu 50 55 60

Gln Phe Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His 65 70 75 80

5

Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys 85 90 95

Gly Leu Pro Ser Ser Ile Glu Lys Thr Ile Ser Lys Ala Lys 100 105 110

<210> 8

<211> 110

<212> PRT

<213> Artificial Sequence

<220>

<400> 8

Ala Pro Glu Leu Leu Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys 1 5 10 15

Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val 20 25 30

Val Val Asp Val Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr
35 40 45

Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu
50 55 60

Gln Tyr Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His
65 70 75 80

Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys 85 90 95

Gly Leu Pro Ser Ser Ile Glu Lys Thr Ile Ser Lys Ala Lys
100 105 110

<210> 9

<211> 109

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Mutated
 antibody

<400> 0

Ala Pro Pro Val Ala Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro
1 5 10 15

Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val 20 25 30

Val Asp Val Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr Val 35 40 45

Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln 50 55 60

Tyr Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln 65 70 75 80

Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Ala 85 90 95

Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Ala Lys 100 105

<210> 10

<211> 110

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Mutated
 antibody

<400> 10

Ala Pro Pro Val Ala Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys 1 5 10 15

Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val 20 25 30

Val Val Asp Val Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr 35 40 45

Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu 50 55 60

Gln Tyr Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His
65 70 75 80

Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys 85 90 95

Ala Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Ala Lys 100 105 110

```
<210> 11
```

<211> 109

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Mutated
 antibody

<400> 11

Ala Pro Pro Val Ala Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro 1 5 10 15

Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val 20 25 30

Val Asp Val Ser Gln Glu Asp Pro Glu Val Gln Phe Asn Trp Tyr Val
35 40 45

Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln
50 55 60

Phe Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln 65 70 75 80

Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Gly 85 90 95

Leu Pro Ser Ser Ile Glu Lys Thr Ile Ser Lys Ala Lys 100 105

<210> 12

<211> 110

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Mutated
 antibody

<400> 12

Ala Pro Pro Val Ala Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys 1 5 10 15

Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val 20 25 30

Val Val Asp Val Ser Gln Glu Asp Pro Glu Val Gln Phe Asn Trp Tyr 35 40 45

Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu 50 55 60

Gln Phe Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His
65 70 75 80

Gln P	Asp	Trp	Leu	Asn 85	Gly	Lys	Glu	Tyr	Lys 90	Cys	Lys	Val	Ser	Asn 95	Lys	
Gly I	Leu	Pro	Ser 100	Ser	Ile	Glu	Lys	Thr 105	Ile	Ser	Lys	Ala	Lys 110			
<210>	. 13	1												-		
<211><211><212><212><213>	> 26 > DN	i A	.cial	L Sec	quenc	ce										
<220> <223>	> De			on of eotic		cific	cial	Seqı	ience	e:						
<400> ggatg			acto	gagg	gg ca	accto	J									26
<210><211><211><212><213>	29 DN	A	cial	. Sec	quenc	ce										
<220> <223>	De			on of eotic		ific	cial	Sequ	ience	:						
<400> tgtcc			ccct	ggta	ic co	cacç	ıggt									29
<210><211><211><212><213>	28 DN	A	cial	. Sec	luenc	e										
<220> <223>	De			on of otid		ific	ial	Sequ	ience	:						
<400> gagcc			ctct	agac	a cc	ctcc	ct									28
<210> <211> <212> <213>	36 DN	A	cial	Seq	uenc	e										
<220> <223>	De			n of		ific	ial	Sequ	ence	:						

totocaacaa aggootooog tootocatog agaaaa	36
<210> 17 <211> 36 <212> DNA <213> Artificial Sequence	
<220> <223> Description of Artificial Sequence: Oligonucleotide	
<400> 17 ttttctcgat ggaggacggg aggcctttgt tggaga	36
<210> 18 <211> 36 <212> DNA <213> Artificial Sequence	
<220> <223> Description of Artificial Sequence: Oligonucleotide	
<400> 18 tecteageae etceagtege ggggggaeeg teagte	36
<210> 19 <211> 33 <212> DNA <213> Artificial Sequence	
<220> <223> Description of Artificial Sequence: Oligonucleotide	
<400> 19 gactgacggt cccgcgactg gaggtgctga gga	33
<210> 20 <211> 26 <212> DNA <213> Artificial Sequence	
<220> <223> Description of Artificial Sequence: Oligonucleotide	
<400> 20 caggtgccct cgagtagcct gcatcc	26

.

<210> 21 <211> 36 <212> DNA <213> Artificial Sequence	
<220> <223> Description of Artificial Sequence: Oligonucleotide	_
<400> 21 tccacaggtg tccactccca ggtgcatcta cagcag	36
<210> 22 <211> 37 <212> DNA <213> Artificial Sequence	
<220> <223> Description of Artificial Sequence: Oligonucleotide	
<400> 22 gaggttgtaa ggactcacct gaggagacgg tgaccgt	37
<210> 23 <211> 36 <212> DNA <213> Artificial Sequence	
<220> <223> Description of Artificial Sequence: Oligonucleotide	
<400> 23 tccacaggtg tccactccga catccagatg acccag	36
<210> 24 <211> 37 <212> DNA <213> Artificial Sequence	
<220> <223> Description of Artificial Sequence: Oligonucleotide	
<400> 24 gaggttgtaa ggactcacgt ttgatctcca gcttggt	37
<210> 25 <211> 20 <212> DNA <213> Artificial Sequence	

```
<220>
<223> Description of Artificial Sequence: Primer
<400> 25
ggagtggaca cctgtggaga
                                                                    20
<210> 26
<211> 20
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Primer
<400> 26
gtgagtcctt acaacctctc
                                                                    20
<210> 27
<211> 6
<212> PRT
<213> Homo sapiens
<400> 27
Glu Leu Leu Gly Gly Pro
```

1